

Husbands also does not describe ODV base as a crystalline material but rather describes a hydrogenation reaction to produce a solid that was then converted to a fumarate salt. Applicants enclose herewith a true and accurate copy (signed and witnessed) of two pages from Dr. Morris Husband's notebook (Attachment A) in which Dr. Husbands recorded the process of Example 26 and in particular describes that "the p-benzyloxy-analog of venlafaxine (referred to as 109-1) was dissolved in 100 ml of EtOH and added to 1.0 g of 10% Pd/C moistened with ethanol. Cyclohexa-1,4-diene (5ml) was added and the mixture stirred for about 90 minutes. The catalyst was filtered and the solvent removed. A glass was obtained. Wt 800 mg." Thus, Husbands does not teach a crystalline form of ODV base.

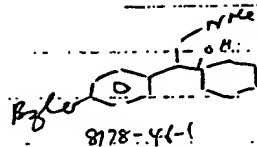
In view of the foregoing, Applicants respectfully maintain that neither Jerussi nor Husbands teach or suggest the invention of Claims 1 and 2 and this rejection should be withdrawn.



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Purification of



5-38-1508-007

EtAc / MeOH 15705

the system

Fr	Vol	Sol	Res	Fr	Vol	Sol	Res
1	500	EtAc	rough	11	200	200	
2	300	"		12	"	"	
3	125	"		13	"	"	
4	200	"		14	"	"	
5	"	"		15	"	"	
6	"	"		16	"	30%	
7	"	"		17	"	"	
8	"	"		18	"	"	
9	"	"					
10	"	"					

109-1

TLC was done on most fractions. The product was present in fractions 10-16. Wt ~ 2 gm.

A small portion was converted to hydrochloride as usual

109-2

Wt:

OBSERVED

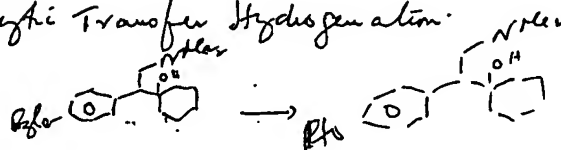
READ

SIGNED

A/Kao

Exp. 130

Catalytic Transfer Hydrogenation



~1.1 gm 109-1 was dissolved in 100 ml EtOH and added to 1.0 gm 10% Pd/C maintained with

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110

ethanol. ~~St~~ cyclohexa-1,4-diene (5 ml) was
added and the mixture stirred for ~ 90 mins.
The catalyst was filtered and the solvent removed.
A glass was obtained.

7970-110-1

AN 800 mg.

7970-110-2

OBSERVED _____
READ W. Kan
SIGNED _____

W. Kan

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